Appl. No. :

10/035,855

Filed

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December 26, 2001

#### REMARKS

Upon entry of the foregoing amendments, the specification has been amended to remove the recitation of URLs. No new matter has been added by the amendments to the specification.

The claims have been amended as set forth above. Claims 22-26 are pending. Claim 27 has been cancelled. Claim 22 has been amended to remove reference to Figure 20 and to recite "specifically binds," which phrase was included in cancelled Claim 27. Also, as discussed more fully below, Claims 22-26 have been amended to recite "isolated." No new matter is added by the amendments and the claims are fully supported by the specification as originally filed.

Applicants respond below to the specific rejections raised by the PTO in the Office Action mailed September 13, 2004.

## **Information Disclosure Statement**

The Examiner asserts that the previously-filed information disclosure statement fails to comply with 37 C.F.R. § 1.98(a)(2). The Examiner notes that the Blast results are not true publications with a publication date, and therefore, are not fully in compliance with 37 C.F.R. § 1.97.

Respectfully, Applicants disagree. The Blast results are true publications with a publication date or other information consistent with the duty of disclosure and § 1.98(a)(2). The contents of the previously filed IDS, which has been objected to, satisfied the requirements of § 1.98(a)(2) because the IDS included a legible copy of each publication or that portion which caused it to be listed and all other information or that portion which caused it to be listed. In particular, the previously filed Blast results were legible, provided a comparison of a claimed sequence to another sequence, and showed the relevant information for the other sequence.

Nonetheless, for the convenience of the Examiner, more detailed information is submitted as Exhibit 1. The resubmitted results include more detailed information regarding the cited sequences. These Blast results are resubmitted before the mailing date of any of a final action under § 1.113, a notice of allowance under § 1.311, or an action that otherwise closes prosecution in the application. Applicants believe that no fee is due because of compliance with §§ 1.97-1.98. However, if a fee is due to ensure consideration of the submitted Blast results, for example, the fee under § 1.17(p), the Patent Office is authorized to charge the fee to Deposit Account No. 11-1410.

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## **Specification**

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The Examiner states that the specification should be reviewed for the recitation of improper hyperlinks, and that all such recitations should be deleted or amended. Applicants have amended the specification to address the Examiner's concern. In particular, Applicants have replaced the hyperlinks with text that describes the location of the websites. The amended text no longer constitutes browser executable code.

## Rejection under 35 U.S.C. §101 - Statutory Subject Matter

The Examiner rejects Claims 22, 25 and 27 under 35 U.S.C. § 101 because allegedly the claimed invention is directed to non-statutory subject matter. The Examiner asserts that the "recitation of 'an antibody' encompasses all naturally occurring antibodies, and antibody fragments thereof, to the naturally occurring polypeptide of SEQ ID NO:45; thereby not involving the hand of man to isolate or purify the antibodies, or fragments thereof." The Examiner suggests amending the claim to recite "isolated" and "purified" antibody.

Claims 22-26 have been amended to recite "isolated." Therefore, reconsideration and withdrawal of the instant rejection is respectfully requested.

### Rejection under 35 U.S.C. §101 - Utility

The Examiner rejects Claims 22-27 as allegedly not being supported by a specific and substantial asserted utility, or a well established utility. The Examiner notes that the claims are directed to antibodies that bind to the polypeptide of SEQ ID NO:45, referred to in the specification as PRO4405. The Examiner argues that utility of the claimed antibodies depends upon the whether or not the polypeptide to which the antibodies bind have utility and enablement. According to the Examiner the PRO4405 polypeptides, to which the claimed antibodies bind, do not have utility.

According to the Utility Examination Guidelines ("Utility Guidelines"), 66 Fed. Reg. 1092 (2001) an invention complies with the utility requirement of 35 U.S.C. § 101, if it has at least one asserted "specific, substantial, and credible utility" or a "well-established utility." A utility is "specific" when it is particular to the subject matter claimed. With regard to substantial utility, "[a]ny reasonable use that an applicant has identified for the invention that can be viewed

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as providing a public benefit should be accepted as sufficient, at least with regard to defining a 'substantial' utility." (M.P.E.P. 2107.01). "Credibility is assessed from the perspective of one of ordinary skill in the art in view of the disclosure and any other evidence of record ... that is probative of the Applicant's assertions." (M.P.E.P. 2107 II(B)(1)(ii)). Such standard is presumptively satisfied unless the logic underlying the assertion is seriously flawed, or if the facts upon which the assertion is based are inconsistent with the logic underlying the assertion (Revised Interim Utility Guidelines Training Materials, 1999).

Respectfully, the claimed antibodies bind to polypeptides that have a specific, substantial and credible utility. The utility for the polypeptides is set forth in Example 36 of the specification at page 166. Example 36 describes a chondrocyte redifferentiation assay (Assay #110). This assay shows that certain polypeptides act to induce redifferentiation of chondrocytes, and therefore, are useful for the treatment of various bone and/or cartilage disorders such as, for example, sports injuries and arthritis. As mentioned in Example 36, PRO4405 is one of polypeptides that tested positive in the chondrocyte redifferentiation assay.

The ability to induce chondrocyte redifferentiation is specific or particular to the PRO4405 polypeptides, and is not an ability common to all peptides generally. Also, the utility is substantial as treatment of bone and/or cartilage disorders provides a public benefit. Finally, one of ordinary skill in the art would recognize that the scientific assay results of Example 36 support the credibility of the utility assertion.

Therefore, the claimed antibodies have utility because the polypeptides to which they bind have a specific, substantial and credible utility.

For the reasons discussed above, Applicants respectfully request reconsideration and withdrawal of the instant rejection under 35 U.S.C. § 101.

# Rejections under 35 U.S.C. §112, first paragraph – Enablement

The Examiner rejected Claims 22-27 under 35 U.S.C. § 112, first paragraph. According to the Examiner, because the claimed invention is not supported by either a substantial asserted utility or a well established utility, on of skill in the art would not know how to use the invention.

Applicants submit that in the above discussion of the utility rejection under 35 U.S.C. § 101, Applicants have established a substantial, specific, and credible utility for the claimed antibodies. Specifically, the claimed antibodies and the polypeptides to which they bind the have

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Filed: December 26, 2001

utility for inducing chondrocyte redifferentiation, which demonstrates useful application for various bone and/or cartilage disorders such as, for example, sports injuries and arthritis. The specification teaches how to make and use the polypeptides and the claimed antibodies. Based upon that teaching and the above-established utility for the claimed subject matter, one skilled in the art would know how to make and use the claimed subject matter.

Therefore, Applicants therefore request that the Examiner reconsider and withdraw the enablement rejection under 35 U.S.C. § 112, first paragraph, based on a lack of utility.

## Rejections under 35 U.S.C. §112, second paragraph - Indefiniteness

The Examiner has rejected Claims 22-27 under 35 U.S.C. § 112, second paragraph, as being indefinite. The Examiner objects to the phrase "binds to" in Claim 22 and to the phrase "specifically binds to" in Claim 27 arguing that neither the art nor the specification provides a clear definition for, or distinction between the two phrases. Thus, according to the Examiner, the metes and bounds of the claims cannot be determined by one of skill in the art.

As set forth above, Claim 27 has been cancelled and Claim 22 has been amended to recite "specifically binds." Applicants submit that the term "specifically binds" has a well established meaning – for example, it refers to the binding of an antibody to a particular polypeptide, where the antibody does not substantially bind to any other polypeptide. One of skill in the art would readily understand the language of the claims to mean that the claimed antibodies bind to specifically defined polypeptides (in this case the polypeptides of SEQ ID NO:45) but do not substantially bind to any other polypeptides. In view of the amendments to the claims, and since claim terms should be given their ordinary, art-recognized meaning, Applicants request reconsideration and withdrawal of the instant rejection.

### Conclusion

The present application is believed to be in condition for allowance, and an early action to that effect is respectfully solicited. Applicants invite the Examiner to call the undersigned if any issues may be resolved through a telephonic conversation.

Please charge any additional fees, including any fees for additional extension of time, or credit overpayment to Deposit Account No. 11-1410.

Appl. No. Filed

10/035,855

December 26, 2001

Respectfully submitted,

KNOBBE, MARTENS, OLSON & BEAR, LLP

Dated: 12/10/04

By:

Marc T. Morley

Registration No. 52,051 Attorney of Record Customer No. 30,313

(619) 235-8550

S:\DOCS\MTM\MTM-7314.DOC 112304

```
###
Tue Jan 8 09:17:51 2002 [BLASTN 2.2.1 [Jul-12-2001], NCBI]
Repeats masked (summary below)
/home/ruby/va/Molbio/carpenda/tempids/ss.DNA84920 (2395 bp)
       /home/ruby/va/Molbio/carpenda/tempids/ss.DNA84920
Database: gen (16,229,280 seqs, 16,995,651,507 bp) Jan 1, 2002
   Locus list: hum -est (1,803,435 seqs, 6,559,376,613 bp)
Matrix: blastn matrix:1 -3, T: 0, A: 40, X1: 6, X2: 15, S1: 12, S2: 20, eval: 10.
Gap Penalties: Existence: 5, Extension: 2
Sequences producing High-scoring Segment Pairs:
                                                     Frame Score Match Pct E-val
  1 P_AAA96345 cDNA encoding a novel polypeptide design
                                                         +
                                                             2395
                                                                   2395 100
                                                                             . 0.0
  2 P AAD02923 Human PRO4405 cDNA (DNA84920-2614). cDN
                                                             2395
                                                                   2395 100
                                                                              0.0
  3 P AAF92127 Human PRO4405 cDNA.
                                                                   2395 100
                                                             2395
                                                                              0.0
  4 P AAC91490 Human PRO4405 cDNA.
                                                             2395
                                                                   2395 100
                                                                              0.0
  5 A\overline{X}089946
               Sequence 7 from Patent WO0116319. DNA,
                                                          + 2395
                                                                              0:0
                                                                   2395 100
  6 AX092408
               Sequence 139 from Patent WO0116318. DNA
                                                             2395
                                                                   2395 100
                                                                              0.0
  7 AX055478
               Sequence 108 from Patent WO0073452. DNA
                                                                              0.0
                                                             2395
GenBank (Release 143, aug 2004)
2395 100
P AAA96345
            cDNA encoding a novel polypeptide designated PRO4405. 395 bp,
          cDNA, PAT 08-FEB-2001
            P AAA96345
ACCESSION
KEYWORDS
            GENESEQ; Secreted protein; transmembrane protein; PRO1484; PRO4334;
          PRO1122; PRO1889; PRO1890; PRO1887; PRO1785; PRO4353; PRO4357;
          PRO4405; PRO4356; PRO4352; PRO4380; PRO4354; PRO4408; PRO5737;
          PRO4425; PRO5990; PRO6030; PRO4424; PRO4422; PRO4430; PRO4499;
          tumour; obesity; diabetes; insulinemia; kidney disorder; Bergers
          disease; nephropathy; Schonlein-Henoch purpura; celiac disease;
          dermatitis herpetiformis; Crohns disease; patent; patentdb
          (v200423, 04-NOV-2004).
SOURCE
            Homo sapiens.
  ORGANISM
            Homo sapiens.
REFERENCE
               (bases 1 to 2395)
  AUTHORS
            Desnoyers, L., Eaton, D.L., Goddard, A., Godowski, P.J.,
          Gurney, A.L., Pan, J. Stewart, T.A., Watanabe, C.K.,
                                                              Wood, W.I.,
          Zhang, Z.
  TITLE
            Novel secreted and transmembrane polypeptides useful for diagnosing
          tumor in a mammal, for identifying agonists and antagonists of the
          polypeptide and for therapeutic use.
            Patent: WO200056889-A2; Filing Date: 01-MAR-2000; 2000WO-US005601;
  JOURNAL
          Publication Date: 28-SEP-2000; Priority: 23-MAR-1999;
                                        99US-0125778P. 24-MAR-1999;
          99US-0125774P. 23-MAR-1999;
          99US-0125826P. 31-MAR-1999;
                                         99US-0127035P. 05-APR-1999;
          99US-0127706P. 21-APR-1999;
                                         99US-0130359P. 27-APR-1999;
                                        99US-0131272P. 27-APR-1999;
          99US-0131270P. 27-APR-1999;
          99US-0131291P. 04-MAY-1999;
                                         99US-0132371P. 04-MAY-1999;
          99US-0132379P. 04-MAY-1999;
                                        99US-0132383P. 25-MAY-1999;
          99US-0135750P. 08-JUN-1999;
                                        99US-0138166P. 20-JUL-1999;
          99US-0144791P. 03-AUG-1999;
                                         99US-0146970P. 09-DEC-1999;
          99US-0170262P; Assignee: (GETH ) GENENTECH INC; Cross Reference:
          WPI; 2000-628263/60. P-PSDB; AAB18918; Patent Format: Claim 2; Fig
          19; 222pp; English.
COMMENT
            The present sequence encodes a secreted or transmembrane
          polypeptide. The specification describes polypeptides designated
          PRO1484, PRO4334, PRO1122, PRO1889, PRO1890, PRO1887, PRO1785,
          PRO4353, PRO4357, PRO4405, PRO4356, PRO4352, PRO4380, PRO4354,
```

PRO4408, PRO5737, PRO4425, PRO5990, PRO6030, PRO4424, PRO4422, PRO4430 and PRO4499. PRO1889 polypeptide is useful for diagnosing tumour in a mammal. The polypeptides, their agonists and antagonists are useful treating a condition associated with expression or activity of the polypeptide. Conditions treated include obesity, diabetes or hyper-or hypo-insulinemia. The polypeptides are capable of inducing proliferation of mammalian kidney mesangial cells and are therefore useful for treating kidney disorders associated with decreased mesangial cell function such as Bergers disease or other nephropathies associated with Schonlein-Henoch purpura, celiac disease, dermatitis herpetiformis or Crohns disease. The nucleic acids may be used to generate transgenic animals for use in development and screening of therapeutically useful reagents and also for chromosome identification and tissue typing

FEATURES

Location/Qualifiers

CDS

79..1011

sig\_peptide

/\*tag= a 79..180

. . . .

/\*tag= b

605 c

BASE COUNT

656 g 568 t

ORIGIN

2395 100 0.0

P\_AAD02923 Human PRO4405 cDNA (DNA84920-2614). 395 bp, cDNA, PAT 31-MAY-2001 ACCESSION P AAD02923

KEYWORDS

GENESEQ; Human; PRO4405; antiinflammatory; dermatological; immunosuppressive; antirheumatic; antiarthritic; osteopathic; antianaemic; haemostatic; antithyroid; antidiabetic; antiviral; antipsoriatic; antiallergic; antiasthmatic; inhibitor; therapy; systemic lupus erythematosis; spondyloarthropathy; systemic sclerosis; systemic vasculitis; sarcoidosis; idiopathic inflammatory myopathy; Sjogren's syndrome; autoimmune thrombocytopenia; immune-mediated renal disease; hepatitis; demyelinating polyneuropathy; Guillian-Barre syndrome; Whipple's disease; hepatobiliary disease; primary biliary cirrhosis; sclerosing cholangitis; inflammatory bowel disease; gluten-sensitive enteropathy; skin disease; allergic rhinitis; atopic dermatitis; food hypersensitivity; urticaria; eosinophilic pneumonia; hypersensitivity pneumonitis; graft rejection; idiopathic pulmonary fibrosis; graft-versus-host-disease; patent; patentdb (v200423, 04-NOV-2004).

SOURCE

Homo sapiens.

ORGANISM Homo sapiens.

REFERENCE 1 (bases 1 to 2395)

AUTHORS Goddard, A., Godowski, P.J., Gurney, A.L., Hillan, K.J., Tumas, D. Watanabe, C.K., Wood, W.I.

TITLE New PRO polypeptides for treating immune related and inflammatory diseases such as rheumatoid arthritis, systemic vasculitis, asthma, autoimmune hemolytic anemia, and diabetes mellitus.

JOURNAL

Patent: WO200116319-A2; Filing Date: 23-AUG-2000; 2000WO-US023522; Publication Date: 08-MAR-2001; Priority: 31-AUG-1999; 99US-0151733P. 01-SEP-1999; 99WO-US020111. 16-DEC-1999; 99WO-US030095. 18-FEB-2000; 2000WO-US004342. 01-MAR-2000; 2000WO-US005601. 30-MAR-2000; 2000WO-US008439. 17-MAY-2000; 2000WO-US013705. 22-MAY-2000; 2000WO-US014042. 30-MAY-2000; 2000WO-US014941. 05-JUN-2000; 2000US-0209832P; Assignee: (GETH )

GENENTECH INC; Cross Reference: WPI; 2001-226690/23. P-PSDB; AAY72877; Patent Format: Claim 2; Fig 7; 118pp; English.

COMMENT

**JOURNAL** 

The present sequence is a cDNA (DNA84920-2614 clone) encoding PRO4405 protein. PRO protein, its agonist or antagonist or its antibody which are capable of enhancing or inhibiting the proliferation of T-lymphocytes or of increasing the infiltration of inflammatory cells into a tissue are useful in the diagnosis and treatment of immune-related diseases in mammals. The PRO protein is useful for treating systemic lupus erythematosis, rheumatoid arthritis, osteoarthritis, juvenile chronic arthritis, spondyloarthropathy, systemic sclerosis, idiopathic inflammatory myopathy, Sjogren's syndrome, systemic vasculitis, sarcoidosis, autoimmune haemolytic anaemia, autoimmune thrombocytopenia, thyroiditis, diabetes mellitus, immune-mediated renal disease, demyelinating disease of the central or peripheral nervous system, idiopathic demyelinating polyneuropathy, Guillian-Barre syndrome, chronic inflammatory demyelinating polyneuropathy, hepatobiliary disease, infectious or autoimmune chronic active hepatitis, primary biliary cirrhosis, granulomatous hepatitis, sclerosing cholangitis, inflammatory bowel disease, gluten-sensitive enteropathy, Whipple's disease, autoimmune or immune-mediated skin diseases such as bullous skin disease, erythema multiforme and contact dermatitis, psoriasis, allergic diseases such as asthma, allergic rhinitis, atopic dermatitis, food hypersensitivity and urticaria, immunologic diseases of the lung such as eosinophilic pneumonias, idiopathic pulmonary fibrosis, hyper- sensitivity pneumonitis, transplantation associated diseases such as graft rejection or graft-versus-host-disease

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FEATURES
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                     79..180
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BASE COUNT
                566 a
                         605 c
                                            568 t
                                   656 g
ORIGIN
2395 100
P_AAF92127 Human PRO4405 cDNA. 395 bp, cDNA, PAT 15-MAY-2001
ACCESSION
            P AAF92127
KEYWORDS
            GENESEQ; Human; PRO protein; mapping; patent; patentdb (v200423,
          04-NOV-2004).
SOURCE
            Homo sapiens.
  ORGANISM Homo sapiens.
REFERENCE
              (bases 1 to 2395)
  AUTHORS
            Eaton, D.L., Filvaroff, E., Gerritsen, M.E., Goddard, A.,
          Godowski, P.J. Grimaldi, C.J., Gurney, A.L., Watanabe, C.K.,
            Eighty four nucleic acids encoding PRO polypeptides, useful in
  TITLE
          molecular biology, including use as hybridization probes, and in
          chromosome and gene mapping.
```

Publication Date: 08-MAR-2001; Priority: 01-SEP-1999; 99WO-US020111. 15-SEP-1999; 99WO-US021090. 07-DEC-1999;

Patent: WO200116318-A2; Filing Date: 24-AUG-2000; 2000WO-US023328;

```
99US-0169495P. 09-DEC-1999;
                                        99US-0170262P. 11-JAN-2000;
          2000US-0175481P. 18-FEB-2000; 2000WO-US004341. 18-FEB-2000;
          2000WO-US004342. 22-FEB-2000; 2000WO-US004414. 01-MAR-2000;
          2000WO-US005601. 03-MAR-2000; 2000US-0187202P. 21-MAR-2000;
          2000US-0191007P. 30-MAR-2000; 2000WO-US008439. 25-APR-2000;
          2000US-0199397P. 22-MAY-2000; 2000WO-US014042. 05-JUN-2000;
          2000US-0209832P; Assignee: (GETH ) GENENTECH INC; Cross Reference:
          WPI; 2001-183260/18. P-PSDB; AAB87595; Patent Format: Claim 2; Fig
          139; 278pp; English.
COMMENT
            The present sequence is the coding sequence for a human PRO
          polypeptide (secreted and transmembrane). The PRO protein, and PRO
          agonists, PRO antagonists or anti-PRO antibodies are useful for
          preparation of a medicament useful in the treatment of a condition
          which is responsive to the PRO protein, agonists, antagonists or
          anti-PRO antibodies. The PRO protein may also be employed as
          molecular weight markers for protein electrophoresis. The PRO
          coding sequence has applications in molecular biology, including
          use as hybridisation probes, and in chromosome and gene mapping
FEATURES
                     Location/Qualifiers
BASE COUNT
                         605 c
                566 a
                                  656 g
                                           568 t
2395 100
           0.0
P AAC91490 Human PRO4405 cDNA. 395 bp, cDNA, PAT 21-MAR-2001
ACCESSION
            P AAC91490
            GENESEQ; Human; PRO; antiinflammatory; dermatological;
KEYWORDS
          antiarthritic; antirheumatic; cardiant; antianaemic;
          immunosuppressive; antithyroid; antidiabetic; nootropic;
          neuroprotective; hepatotropic; virucide; antiallergic;
          antiasthmatic; immune related disorder; hepatobiliary disease;
          autoimmune disease; allergy; patent; patentdb (v200423,
          04-NOV-2004).
          Homo sapiens.
  ORGANISM Homo sapiens.
REFERENCE
           1
               (bases 1 to 2395)
 AUTHORS
           Ashkenazi, A.J., Baker, K.P., Chan, B., Goddard, A., Godowski, P.J.
         Gurney, A.L., Hebert, C., Henzel, W., Kabakoff, R.C., Shelton, D.L.,
          Tumas, D. Watanabe, C.K., Wood, W.I.
 TITLE
            Thirty three nucleic acids encoding PRO polypeptides which are
         useful in the diagnosis and treatment of immune related disorders,
          e.g. systemic lupus erythematosis, rheumatoid arthritis,
         osteoarthritis, thyroiditis and diabetes mellitus.
 JOURNAL
            Patent: WO200073452-A2; Filing Date: 02-JUN-2000; 2000WO-US015264;
         Publication Date: 07-DEC-2000; Priority: 02-JUN-1999;
          99WO-US012252. 20-JUL-1999;
                                       99US-0144732P. 20-JUL-1999;
          99US-0144758P. 28-JUL-1999;
                                        99US-0146222P. 01-SEP-1999;
         99WO-US020111. 15-SEP-1999;
                                        99WO-US021090. 15-SEP-1999;
         99WO-US021547. 29-OCT-1999;
                                        99US-0162506P. 30-NOV-1999;
         99WO-US028313. 01-DEC-1999; 99WO-US028634. 02-DEC-1999;
         99WO-US028551. 02-DEC-1999;
                                        99WO-US028565. 09-DEC-1999;
         99US-0170262P. 20-DEC-1999;
                                        99WO-US030911. 05-JAN-2000;
         2000WO-US000219. 06-JAN-2000; 2000WO-US000376. 11-FEB-2000;
         2000WO-US003565. 18-FEB-2000; 2000WO-US004341. 18-FEB-2000;
         2000WO-US004342. 22-FEB-2000; 2000WO-US004414. 24-FEB-2000;
         2000WO-US004914. 24-FEB-2000; 2000WO-US005004. 01-MAR-2000;
         2000WO-US005601. 02-MAR-2000; 2000WO-US005841. 03-MAR-2000;
         2000US-0187202P. 15-MAR-2000; 2000WO-US006884. 20-MAR-2000;
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ORIGIN

SOURCE

2000WO-US007377. 21-MAR-2000; 2000WO-US007532. 30-MAR-2000; 2000WO-US008439. 17-MAY-2000; 2000WO-US013705. 22-MAY-2000; 2000WO-US014042; Assignee: (GETH ) GENENTECH INC; Cross Reference: WPI; 2001-025253/03. P-PSDB; AAB50931; Patent Format: Claim 48; Fig 59; 218pp; English.

COMMENT

The present sequence is one of thirty three nucleic acids encoding polypeptides. The PRO polypeptides, anti-PRO antibodies, agonists and antagonists are useful for treating and diagnosing immune related disorders such as systemic lupus erythematosus, rheumatoid arthritis, osteoarthritis, juvenile chronic arthritis, spondyloarthropathies, systemic sclerosis, idiopathic inflammatory myopathies, Sjogren's syndrome, systemic vasculitis, sarcoidosis, autoimmune haemolytic anaemia, autoimmune thrombocytopaenia, thyroiditis, diabetes mellitus, immune-mediated renal disease, demyelinating diseases of the central and peripheral nervous systems (such as multiple sclerosis, idiopathic demyelinating polyneuropathy or Guillain-Barre syndrome, and chronic inflammatory demyelinating polyneuropathy), hepatobiliary diseases (such as infectious, autoimmune chronic active hepatitis, primary biliary cirrhosis, granulomatous hepatitis and sclerosing cholangitis), inflammatory bowel disease, gluten-sensitive enteropathy and Whipple's disease, autoimmune or immune-mediated skin diseases (such as bullous skin diseases, erythema multiforme, contact dermatitis, psoriasis), allergic diseases such as asthma, allergic rhinitis, atopic dermatitis, food hypersensitivity and urticaria), immunological diseases of the lung. (such as eosinophilic pneumonias, idiopathic pulmonary fibrosis and hypersensitivity pneumonitis), transplantation associated diseases including graft rejection and graft-versus-host diseases

FEATURES

Location/Qualifiers

BASE COUNT 566 a

566 a 605 c 656 g 568 t

ORIGIN

2395 100 0.0

AX089946 Sequence 7 from Patent W00116319. 2395 bp,

DNA, linear, PAT 21-MAR-2001

ACCESSION AX089946

VERSION AX089946.1 GI:13443984

KEYWORDS

SOURCE Homo sapiens (human)

ORGANISM Homo sapiens

Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.

REFERENCE 1

AUTHORS Goddard, A., Godowski, P.J., Gurney, A.L., Hillan, K.J., Tumas, D., Watanabe, C.K. and Wood, W.I.

TITLE Compositions and methods for the treatment of immune related diseases

JOURNAL Patent: WO 0116319-A 7 08-MAR-2001;

Genentech, Inc. (US)

FEATURES Location/Qualifiers

source 1..2395

/organism="Homo sapiens"
/mol\_type="unassigned DNA"
/db\_xref="taxon:9606"

BASE COUNT ORIGIN

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2395 100 . 0.0
AX092408
            Sequence 139 from Patent WO0116318. 2395 bp,
          DNA, linear, PAT 21-MAR-2001
ACCESSION
            AX092408
            AX092408.1 GI:13444518
VERSION
KEYWORDS
SOURCE
            Homo sapiens (human)
  ORGANISM Homo sapiens
          Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
          Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
REFERENCE
  AUTHORS
            Eaton, D.L., Filvaroff, E., Gerritsen, M.E., Goddard, A.,
          Godowski, P.J., Grimaldi, C.J., Gurney, A.L., Watanabe, C.K. and
          Wood, W.I.
  TITLE
            Secreted and transmembrane polypeptides and nucleic acids encoding
          the same
  JOURNAL
            Patent: WO 0116318-A 139 08-MAR-2001;
          Genentech, Inc. (US)
                      Location/Qualifiers
FEATURES
                      1..2395
     source
                    /organism="Homo sapiens"
                    /mol_type="unassigned DNA"
                    /db xref="taxon:9606"
BASE COUNT
ORIGIN
2395 100
AX055478
            Sequence 108 from Patent WO0073452. 2395 bp,
          DNA, linear, PAT 13-JAN-2001
ACCESSION
            AX055478
VERSION
            AX055478.1 GI:12228736
KEYWORDS.
SOURCE
            Homo sapiens (human)
  ORGANISM Homo sapiens
          Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
          Mammalia; Eutheria; Primates; Catarrhini; Hominidae; Homo.
REFERENCE
            Ashkenazi, A.J., Baker, K.P., Chan, B., Goddard, A., Godowski, P.J.,
  AUTHORS
          Gurney, A.L., Hebert, C., Henzel, W., Kabakoff, R.C., Shelton, D.L.,
          Tumas, D., Watanabe, C.K. and Wood, W.I.
  TITLE
            Compositions and methods for the treatment of immune related
          diseases
  JOURNAL
            Patent: WO 0073452-A 108 07-DEC-2000;
          Genentech, Inc. (US)
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ORIGIN
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Mon Jan 7 16:13:00 2002 [BLASTP 2.2.1 [Jul-12-2001], NCBI]
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Matrix: BLOSUM62, T: 11, A: 40, X1: 16, X2: 38, X3: 64, S1: 41, S2: 71, eval: 10.
Gap Penalties: Existence: 11, Extension: 1
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                                                             Score Match Pct E-val
  1 P AAB87595
                 Human PRO4405 - Homo sapiens.
                                                              1617
                                                                     310 100 e-179
  2 P AAY72877
                  Human PRO4405 protein encoded by DNA84920
                                                              1617
                                                                     310 100 e-179
                 Human polypeptide, SEQ ID NO: 2891 - Homo 1617 310 100 e-179 novel polypeptide designated PRO4405 - Ho 1617 310 100 e-179
  3 P AAM93346
  4 P AAB18918
Dayhoff Protein Database (Rel 78, Mar 2004)
P_AAB87595 Human PRO4405 - Homo sapiens.
Length: 310 aa
Accession: P AAB87595;
Species: Homo sapiens.
Keywords: Human; PRO protein; mapping; patent; GENESEQ patentdb.
Patent number: WO200116318-A2.
Publication date: 08-MAR-2001.
Filing date: 24-AUG-2000; 2000WO-US023328.
Priority: 01-SEP-1999; 99WO-US020111. 15-SEP-1999; 99WO-US021090.
       07-DEC-1999; 99US-0169495P. 09-DEC-1999; 99US-0170262P.
      11-JAN-2000; 2000US-0175481P. 18-FEB-2000; 2000WO-US004341.
      18-FEB-2000; 2000WO-US004342. 22-FEB-2000; 2000WO-US004414.
      01-MAR-2000; 2000WO-US005601. 03-MAR-2000; 2000US-0187202P.
      21-MAR-2000; 2000US-0191007P. 30-MAR-2000; 2000WO-US008439.
      25-APR-2000; 2000US-0199397P. 22-MAY-2000; 2000WO-US014042.
      05-JUN-2000; 2000US-0209832P.
Assignee: (GETH ) GENENTECH INC.
Inventors: Eaton DL, Filvaroff E, Gerritsen ME, Goddard A, Godowski PJ;
      Grimaldi CJ, Gurney AL, Watanabe CK, Wood WI;
Cross reference: WPI; 2001-183260/18. N-PSDB; AAF92127.
Title: Eighty four nucleic acids encoding PRO polypeptides, useful in
      molecular biology, including use as hybridization probes, and in
      chromosome and gene mapping.
Patent format: Claim 12; Fig 140; 278pp; English.
Comment: The present sequence is a human PRO polypeptide (secreted and
      transmembrane). The PRO protein, and PRO agonists, PRO antagonists
      or anti-PRO antibodies are useful for preparation of a medicament
      useful in the treatment of a condition which is responsive to the
      PRO protein, agonists, antagonists or anti-PRO antibodies. The PRO
      protein may also be employed as molecular weight markers for
      protein electrophoresis. The PRO coding sequence has applications
      in molecular biology, including use as hybridisation probes, and in
      chromosome and gene mapping
Database: GENESEQ patent database (v200423, 04-NOV-2004).
P AAY72877
            Human PRO4405 protein encoded by DNA84920-2614 cDNA clone -
            Homo sapiens.
Length: 310 aa
Accession: P_AAY72877;
Species: Homo sapiens.
Keywords: Human; PRO4405; antiinflammatory; dermatological;
      immunosuppressive; antirheumatic; antiarthritic; osteopathic;
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antianaemic; haemostatic; antithyroid; antidiabetic; antiviral; antipsoriatic; antiallergic; antiasthmatic; inhibitor; therapy; systemic lupus erythematosis; spondyloarthropathy; systemic sclerosis; systemic vasculitis; sarcoidosis; idiopathic inflammatory myopathy; Sjogren's syndrome; autoimmune thrombocytopenia; immune-mediated renal disease; hepatitis; demyelinating polyneuropathy; Guillian-Barre syndrome; Whipple's disease; hepatobiliary disease; primary biliary cirrhosis; sclerosing cholangitis; inflammatory bowel disease; gluten-sensitive enteropathy; skin disease; allergic rhinitis; atopic dermatitis; food hypersensitivity; urticaria; eosinophilic pneumonia; hypersensitivity pneumonitis; graft rejection; idiopathic pulmonary fibrosis; graft-versus-host-disease; patent; GENESEQ patentdb.

Patent number: WO200116319-A2.

Publication date: 08-MAR-2001.

Filing date: 23-AUG-2000; 2000WO-US023522.

Priority: 31-AUG-1999; 99US-0151733P. 01-SEP-1999; 99WO-US020111.

16-DEC-1999; 99WO-US030095. 18-FEB-2000; 2000WO-US004342.

01-MAR-2000; 2000WO-US005601. 30-MAR-2000; 2000WO-US008439.

17-MAY-2000; 2000WO-US013705. 22-MAY-2000; 2000WO-US014042.

30-MAY-2000; 2000WO-US014941. 05-JUN-2000; 2000US-0209832P.

Assignee: (GETH ) GENENTECH INC.

Cross reference: WPI; 2001-226690/23. N-PSDB; AAD02923.

Title: New PRO polypeptides for treating immune related and inflammatory diseases such as rheumatoid arthritis, systemic vasculitis, asthma, autoimmune hemolytic anemia, and diabetes mellitus.

Patent format: Claim 10; Fig 8; 118pp; English.

Comment: The present sequence is PRO4405 protein encoded by DNA84920-2614 cDNA clone. PRO protein, its agonist or antagonist or its antibody which are capable of enhancing or inhibiting the proliferation of T-lymphocytes or of increasing the infiltration of inflammatory cells into a tissue are useful in the diagnosis and treatment of immune-related diseases in mammals. The PRO protein is useful for treating systemic lupus erythematosis, rheumatoid arthritis, osteoarthritis, juvenile chronic arthritis, spondyloarthropathy, systemic sclerosis, idiopathic inflammatory myopathy, Sjogren's syndrome, systemic vasculitis, sarcoidosis, autoimmune haemolytic anaemia, autoimmune thrombocytopenia, thyroiditis, diabetes mellitus, immune-mediated renal disease, demyelinating disease of the central or peripheral nervous system, idiopathic demyelinating polyneuropathy, Guillian-Barre syndrome, chronic inflammatory demyelinating polyneuropathy, hepatobiliary disease, infectious or autoimmune chronic active hepatitis, primary biliary cirrhosis, granulomatous hepatitis, sclerosing cholangitis, inflammatory bowel disease, gluten-sensitive enteropathy, Whipple's disease, autoimmune or immune-mediated skin diseases such as bullous skin disease, erythema multiforme and contact dermatitis, psoriasis, allergic diseases such as asthma, allergic rhinitis, atopic dermatitis, food hypersensitivity and urticaria, immunologic diseases of the lung such as eosinophilic pneumonias, idiopathic pulmonary fibrosis, hyper- sensitivity pneumonitis, transplantation associated diseases such as graft rejection or graft-versus-host-disease

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58-76/Domain
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278-284/Modified-site
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Database: GENESEQ patent database (v200423, 04-NOV-2004).
P AAM93346 Human polypeptide, SEQ ID NO: 2891 - Homo sapiens.
Length: 975 aa
Accession: P AAM93346;
Species: Homo sapiens.
Keywords: Human; full length cDNA; cDNA synthesis; oligo-capping; patent;
      GENESEQ patentdb.
Patent number: EP1130094-A2.
Publication date: 05-SEP-2001.
Filing date: 07-JUL-2000; 2000EP-00114089.
Priority: 08-JUL-1999; 99JP-00194486. 11-JAN-2000; 2000JP-00118774.
      02-MAY-2000; 2000JP-00183765.
Assignee: (HELI-) HELIX RES INST.
Inventors: Ota T, Nishikawa T, Isogai T, Hayashi K, Ishii S, Kawai Y;
      Wakamatsu A, Sugiyama T, Nagai K, Kojima S, Otsuki T, Koga H;
Cross reference: WPI; 2001-524255/58. N-PSDB; AAK94266.
Title: 830 Primers useful for synthesizing full length cDNA clones and
      their use in genetic manipulation.
Patent format: Claim 8; SEQ ID NO 2891; 1380pp + Sequence Listing; English.
Comment: The invention relates to primers for synthesising full length cDNA
      clones. 830 cDNA molecules encoding a human protein have been
      isolated and nucleotide sequences of 5'- and 3'-ends of the cDNA
      molecules have been determined. Primers for synthesising the full
      length cDNA are useful for clarifying the function of the protein
      encoded by the cDNA. The full length clones were obtained by
      construction of full length enriched cDNA libraries that were
      synthesised by the oligo-capping method. The primers enable the
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production of the full length cDNA easily without any special methods. The present sequence is a polypeptide encoded by a full length human cDNA of the invention. Note: The sequence data for this patent did not form part of the printed specification, but was obtained in CD-ROM format directly from EPO Database: GENESEQ patent database (v200423, 04-NOV-2004). P AAB18918 A novel polypeptide designated PRO4405 - Homo sapiens. Length: 310 aa Accession: P AAB18918; Species: Homo sapiens. Keywords: Secreted protein; transmembrane protein; PRO1484; PRO4334; PRO1122; PRO1889; PRO1890; PRO1887; PRO1785; PRO4353; PRO4357; PRO4405; PRO4356; PRO4352; PRO4380; PRO4354; PRO4408; PRO5737; PRO4425; PRO5990; PRO6030; PRO4424; PRO4422; PRO4430; PRO4499; tumour; obesity; diabetes; insulinemia; kidney disorder; Bergers disease; nephropathy; Schonlein-Henoch purpura; celiac disease; dermatitis herpetiformis; Crohns disease; patent; GENESEQ patentdb. Patent number: WO200056889-A2. Publication date: 28-SEP-2000. Filing date: 01-MAR-2000; 2000WO-US005601. Priority: 23-MAR-1999; 99US-0125774P. 23-MAR-1999; 99US-0125778P. 24-MAR-1999; 99US-0125826P. 31-MAR-1999; 99US-0127035P. 05-APR-1999; 99US-0127706P. 21-APR-1999; 99US-0130359P. 27-APR-1999; 99US-0131270P. 27-APR-1999; 99US-0131272P. 27-APR-1999; 99US-0131291P. 04-MAY-1999; 99US-0132371P. 04-MAY-1999; 99US-0132379P. 04-MAY-1999; 99US-0132383P. 25-MAY-1999; 99US-0135750P. 08-JUN-1999; 99US-0138166P. 20-JUL-1999; 99US-0144791P. 03-AUG-1999; 99US-0146970P. 09-DEC-1999; 99US-0170262P. Assignee: (GETH ) GENENTECH INC. Inventors: Desnoyers L, Eaton DL, Goddard A, Godowski PJ, Gurney AL, Pan J; Stewart TA, Watanabe CK, Wood WI, Zhang Z; Cross reference: WPI; 2000-628263/60. N-PSDB; AAA96345. Title: Novel secreted and transmembrane polypeptides useful for diagnosing tumor in a mammal, for identifying agonists and antagonists of the polypeptide and for therapeutic use. Patent format: Claim 12; Fig 20; 222pp; English. Comment: The present sequence represents a secreted or transmembrane polypeptide. The specification describes polypeptides designated PRO1484, PRO4334, PRO1122, PRO1889, PRO1890, PRO1887, PRO1785, PRO4353, PRO4357, PRO4405, PRO4356, PRO4352, PRO4380, PRO4354, PRO4408, PRO5737, PRO4425, PRO5990, PRO6030, PRO4424, PRO4422, PRO4430 and PRO4499. PRO1889 polypeptide is useful for diagnosing tumour in a mammal. The polypeptides, their agonists and antagonists are useful treating a condition associated with expression or activity of the polypeptide. Conditions treated include obesity, diabetes or hyper-or hypo-insulinemia. The polypeptides are capable of inducing proliferation of mammalian kidney mesangial cells and are therefore useful for treating kidney disorders associated with decreased mesangial cell function such as Bergers disease or other nephropathies associated with Schonlein-Henoch purpura, celiac disease, dermatitis herpetiformis or Crohns disease. The nucleic acids may be used to generate

transgenic animals for use in development and screening of therapeutically useful reagents and also for chromosome

identification and tissue typing

1-34/Peptide /note= signal peptide/ 6-12/Modified-site /note= N-myristoylation site/ 52-58/Modified-site /note= N-myristoylation site/ 56-60/Modified-site /note= N-glycosylation site/ 58-76/Domain /note= transmembrane domain/ 100-106/Modified-site /note= N-myristoylation site/ 125-131/Modified-site /note= N-myristoylation site/ 154-158/Modified-site /note= amidation site/ 194-198/Modified-site /note= N-glycosylation site/ 233-239/Modified-site /note= N-myristoylation site/ 270-276/Modified-site /note= N-myristoylation site/ 275-281/Modified-site /note= N-myristoylation site/ 278-284/Modified-site /note= N-myristoylation site/ Database: GENESEQ patent database (v200423, 04-NOV-2004).